

Guidelines for the Prevention of Infection Following Combat-related Injuries

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MEDICAL
SYMPOSIUM
ISSUE

Combat-related Infections

Guidelines

- Sponsored by the US Army Office of the Surgeon General
- Conference directors
 - COL Duane R. Hospenthal
 - COL John B. Holcomb
 - MAJ Clinton K. Murray

Guidelines

Conference Goals

- Develop clinical practice guidelines to prevent infections associated with traumatic combat injuries

Guidelines

Conference Goals

- Provide overall guidance to US military health care providers, both deployed and in permanent medical treatment facilities, based on the echelon at which care is provided

Guidelines

Conference Goals

- Accompanying evidence based manuscripts providing guidance by anatomical site of wounding
 - Extremity
 - Central Nervous System
 - Head and Neck
 - Thorax and Abdomen
 - Burn

Guidelines

Participants

- US Army specialty consultants (surgical and infectious disease)
- Participation by US Air Force, US Navy, and civilian trauma experts
- Representatives from the infection control and preventive medicine community

Guidelines

Participants

- LTC RC Andersen
- JH Calhoun, MD
- COL LC Cancio
- MAJ KK Chung
- Maj NG Conger
- HK Crouch
- Maj LC D'Avignon
- COL JR Ficke
- LTC RG Hale
- COL DK Hayes
- EF Hirsch, MD
- MAJ JR Hsu
- Col DH Jenkins
- LCDR JJ Keeling
- COL LE Moores
- CDR KN Petersen
- JR Saffle, MD
- JS Solomkin, MD
- CAPT SA Tasker
- AB Valadka, MD
- LTC AR Wiesen
- COL GW Wortmann

Combat-related Infections

Overview

- Historical review
- Current OIF/OEF epidemiology
- Guideline development
- Guidelines



Greeks

Homer's Iliad and Odyssey

- Therapy
 - Mechanical debridement- remove arrow
 - Rinse wound with warm water or wine
 - Cover wound with bandage soaked in wine
 - Apply analgesic
 - Apply styptic herbal drugs

Achilles
bandages
the arm of
Patroclus



Napoleonic Wars

Amputations

- Larrey performed 200 battlefield amputations in a single day without using anesthesia
- One every 7.2 minutes
 - Hip- 15 seconds
 - Shoulder- 11 seconds
- 80% died



Civil War Federal Troops

- No antiseptics used on wounds
- Operated in pus stained white coats
- Wounds explored with unwashed fingers



Civil War Knowledge

- Laudable pus
- Malignant pus
- Overall wound fatality 14.5%



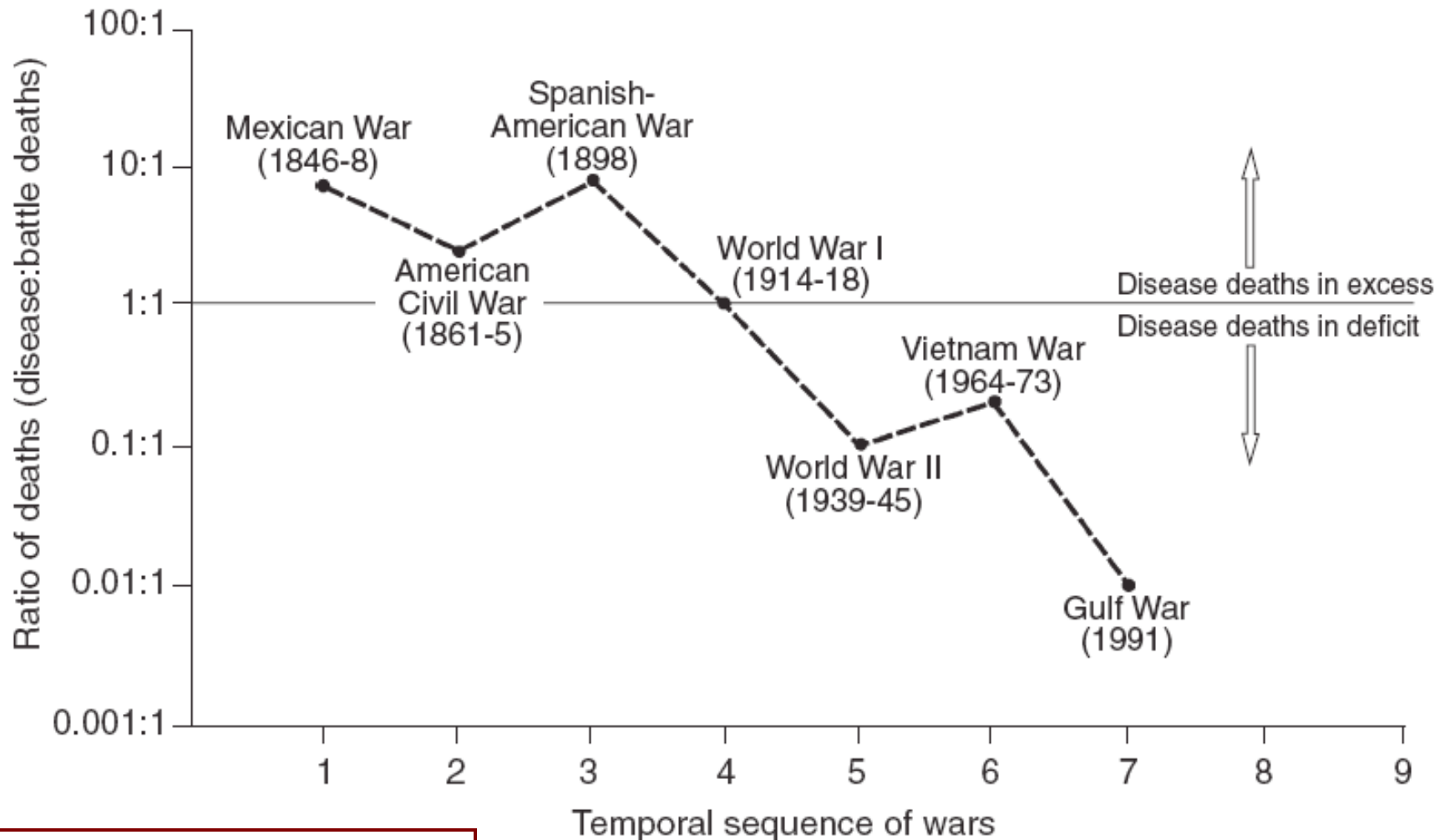
History of Modern ID

Era of Microbiology

- 1862- germ theory (Pasteur)
- 1867- antiseptic surgery (Lister)
- 1881- growth of bacteria on solid media (Koch)
- 1884- gram stain (Gram)



Disease to Battle Deaths Transition



World War I

Modern Surgical Management

- Appropriate surgical management likely led to the disappearance of *Clostridium* associated gas gangrene
 - WWI- 5% incidence with 28% mortality
 - WWII- 1.5% incidence with 15% mortality
 - Korea- 0.08% incidence with no mortality



WWI OR

World War I

Timing of Procedure

- Patients treated
 - Within 1 hour- 10% mortality
 - After 8 hours- 75% mortality



History of Modern ID

Era of Antibiotics

- 1936- Sulfanilamide
- 1942/3- Penicillin

Forrest 1982; Kiehn
1989

Thanks to PENICILLIN
...He Will Come Home!



FROM ORDINARY
MOLD—
*the Greatest Healing
Agent of this War!*

On the green, yellow and blue mold above, called *Penicillium notatum* in the laboratory, grows the miraculous substance first discovered by Professor Alexander Fleming in 1928. Named penicillin by its discoverer, it is the most potent weapon ever developed against many of the deadliest infections known to man. Because research on molds was already a part of Schenley enterprise, Schenley Laboratories were well able to meet the problem of large-scale production of penicillin, when the great need for it arose.

When the thunderous battles of this war have subsided to pages of silent print in a history book, the greatest news event of World War II may well be the discovery and development — not of some vicious secret weapon that destroys — but of a weapon that saves lives. This weapon, of course, is penicillin.

Every day, penicillin is performing some unbelievable act of healing on some far battlefield. Thousands of men will return home who otherwise would not have had a chance. Better still, more and more of this precious drug is now available for civilian use... to save the lives of patients of every age.

A year ago, production of penicillin was difficult, costly. Today, due to specially devised methods of mass-production, in use by Schenley Laboratories, Inc. and the 20 other firms designated by the government to make penicillin, it is available in ever-increasing quantity, at progressively lower cost.

Listen to "THE DOCTOR FIGHTS" starring RAYMOND MASSEY! Tuesday evenings, 8-9 P.M. See your paper for time and station.

SCHENLEY LABORATORIES, INC.
Greenwich, Indiana
Producers of PENICILLIN-Schenley



World War II

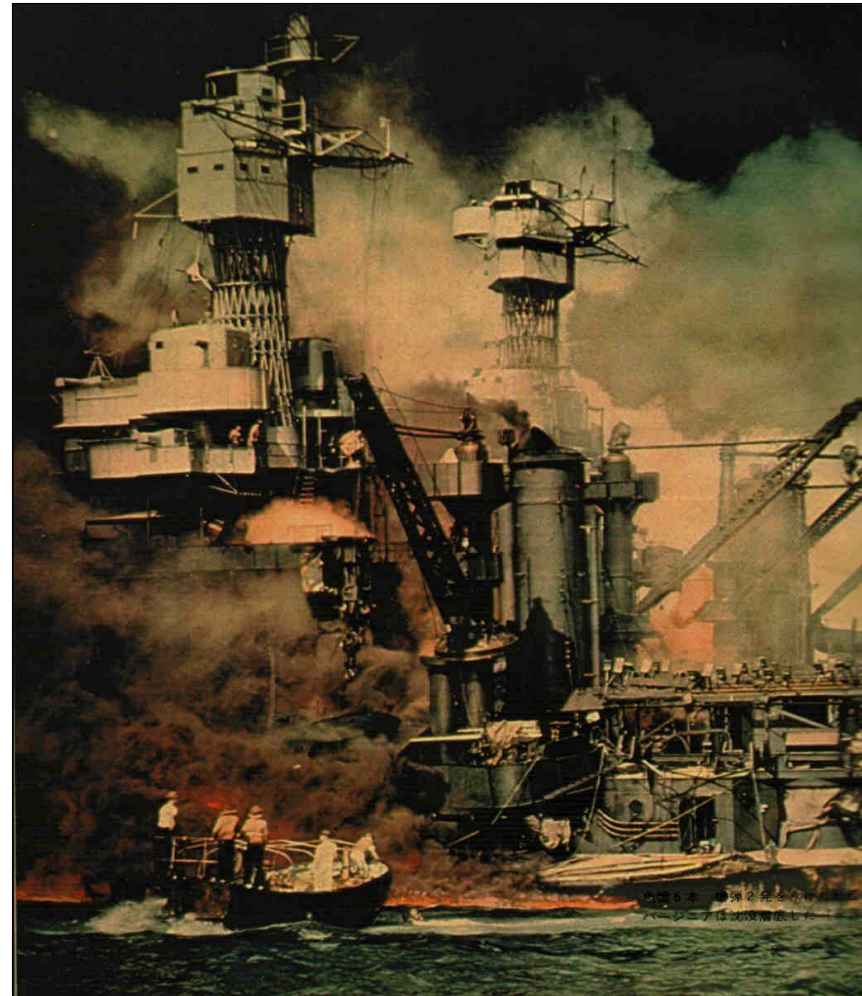
Pearl Harbor

- November of 1941, Dr. John J. Moorhead, chief surgeon for the New York Subway System, came to Honolulu
- Described the techniques for treatment of large soft tissue wounds

World War II

Pearl Harbor

- Only a week later-
Pearl Harbor was
attacked



World War II

Pearl Harbor

- Procedures used
 - Careful debridement
 - Irrigation of wounds
 - Sprinkled in sulfanilamide powder (which was in salt shakers)
 - Left the wounds open and performed a delayed primary closure after three days

World War II

Pearl Harbor

- The immediate reaction “sulfanilamide powder is wonderful”
- The Surgeon General recommended that small packets of sulfanilamide powder be included in the first aid kit of every soldier



World War II

Lessons Learned

- Sulfanilamide powder was dumped in as a lump rather than being sprinkled in so that each grain was separate
- No debride



World War II

Penicillin and Group A Strep

- Eventually systemic penicillin was used during World War II
- It eradicated infections with *S. pyogenes*



World War II

Flap and External fixation

- Applying pedicle graft
- Using Stadler external fixation apparatus



World War II

Nosocomial Transmission

- 5% of wounds were secondarily infected at the time of admission
- 50% were infected after 1 week of hospitalization
- 70-80% were infected after that

Altemeir 1944



MEDICAL CARE

OUT OF EVERY HUNDRED SOLDIERS
RECEIVING MEDICAL CARE
DEATHS IN OUR LAST 2 WARS
WERE

8 DIED



WORLD
WAR I

4 DIED



WORLD
WAR II

Korean War

Antibiotic Era

- Instituted use of penicillin and streptomycin as wound prophylaxis
- Increasingly resistant bacteria were reported from infected war wounds 3-5 days after



Kovaric 1968; Wannamaker
1958

Vietnam War

Antibiotic Era

- During Vietnam broader spectrum of antimicrobial agents were implemented
- Increasingly resistant bacteria were reported in war wounds

Kovaric
1968



Vietnam War

Causes of Death

- Vietnam war- surgical patients 93% (1,162)
 - 43% (494) head injuries
 - 24% (278) hemorrhagic shock
 - **12% (136) septic shock**

Arnold 1978



Vietnam War

Infections

- 4% incidence of wound infections (not including infections after air evacuation)
 - 80% underwent debridement
 - 70% received



Jacob 1989

Vietnam War

Wound cultures 1967-1968-

Japan

Bacteria	% of total positive initial culture (1,153 cultures)
<i>S. aureus</i>	29
<i>P. aeruginosa</i>	18
<i>E. coli</i>	17
Polymicrobial	12
<i>Proteus</i> species	6

Vietnam War

Brooke General Hospital

Bacteria from tissue biopsy	%
<i>P. aeruginosa</i>	47
<i>S. aureus</i>	20
<i>Proteus</i> species	13
<i>Klebsiella</i> <i>Enterobacter</i> group	12
<i>Streptococcus pyogenes</i>	5

Vietnam War

Burn Casualties

- Stabilized in Japan- 106th General Hospital
- Established December 1965



FIGURE 23.—General Heaton observed patient care at an Army hospital in South Vietnam during a visit in November 1967. (Walter Reed Army Institute of Research photograph.)

Operation Just Cause

Wound Infections

- 37 open fractures- 9 infected
 - *CNS*
 - *P. aeruginosa*
- Surgery in the US vs Panama was associated with more infections

Jacob 1992



Gulf War I

Wound Infections

- No well described study assessing infections associated with trauma



Somalia

Wound Infections

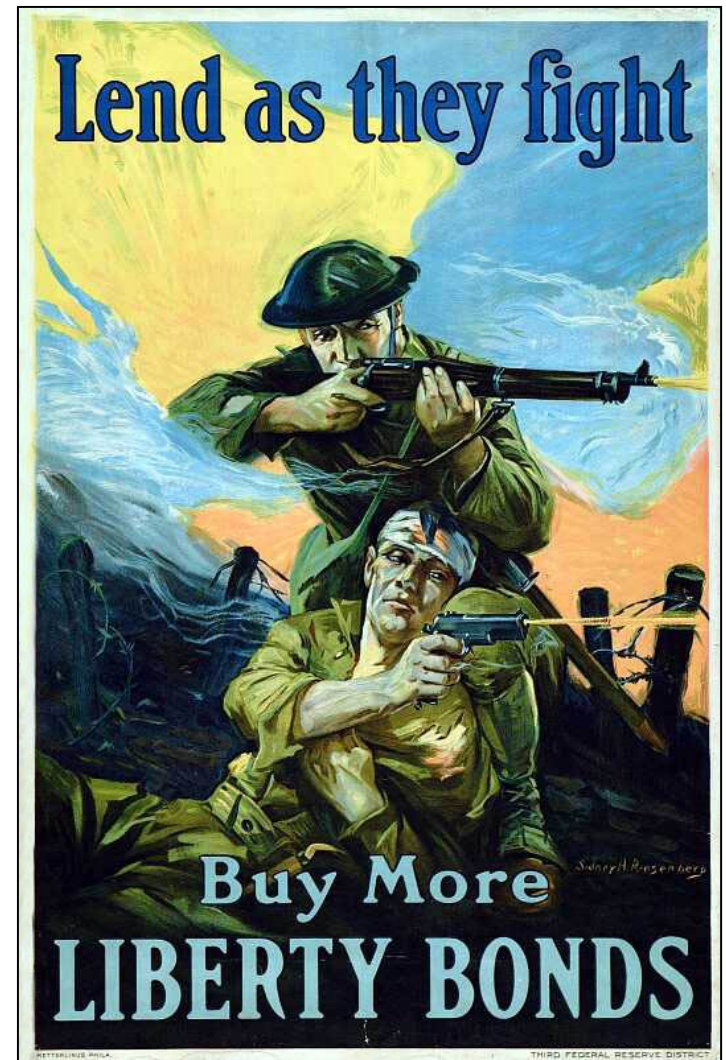
- 11 of 58 wounded in action infected
- Bacteria identified
 - Polymicrobial
 - *Pseudomonas*



Combat-related Infections

Overview

- Historical review
- Current OIF/OEF epidemiology
- Guideline development
- Guidelines





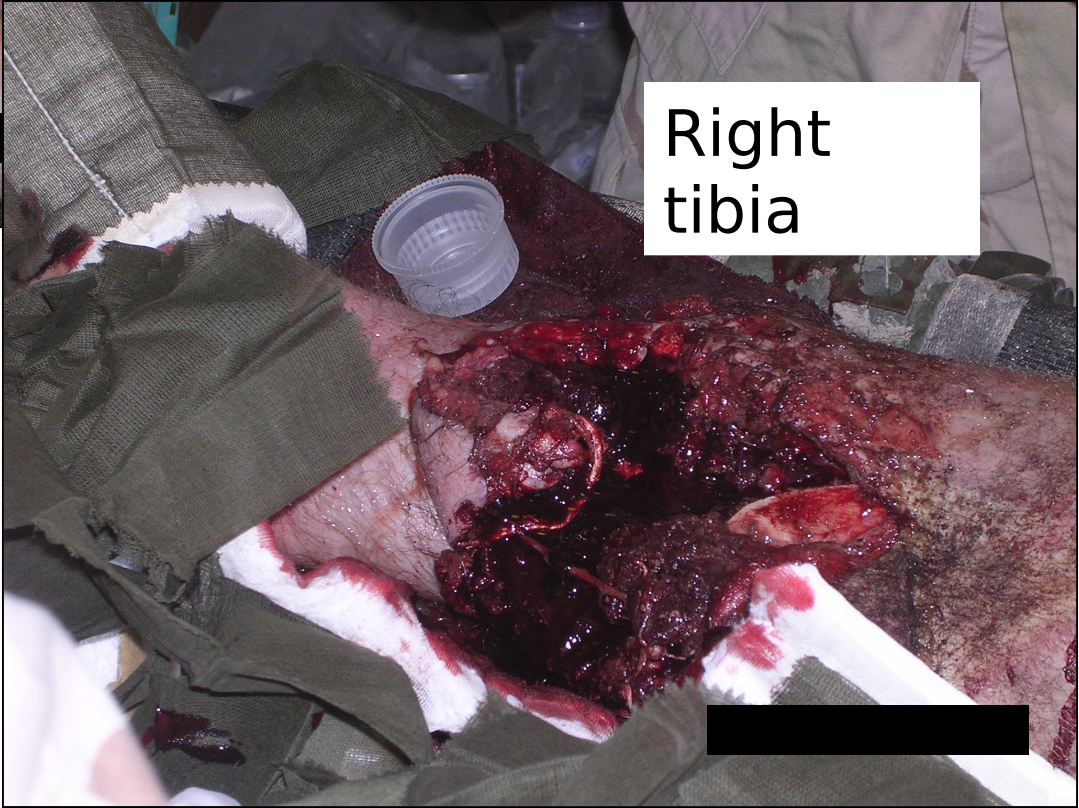




Combat-related Infections

Percentage of Injury

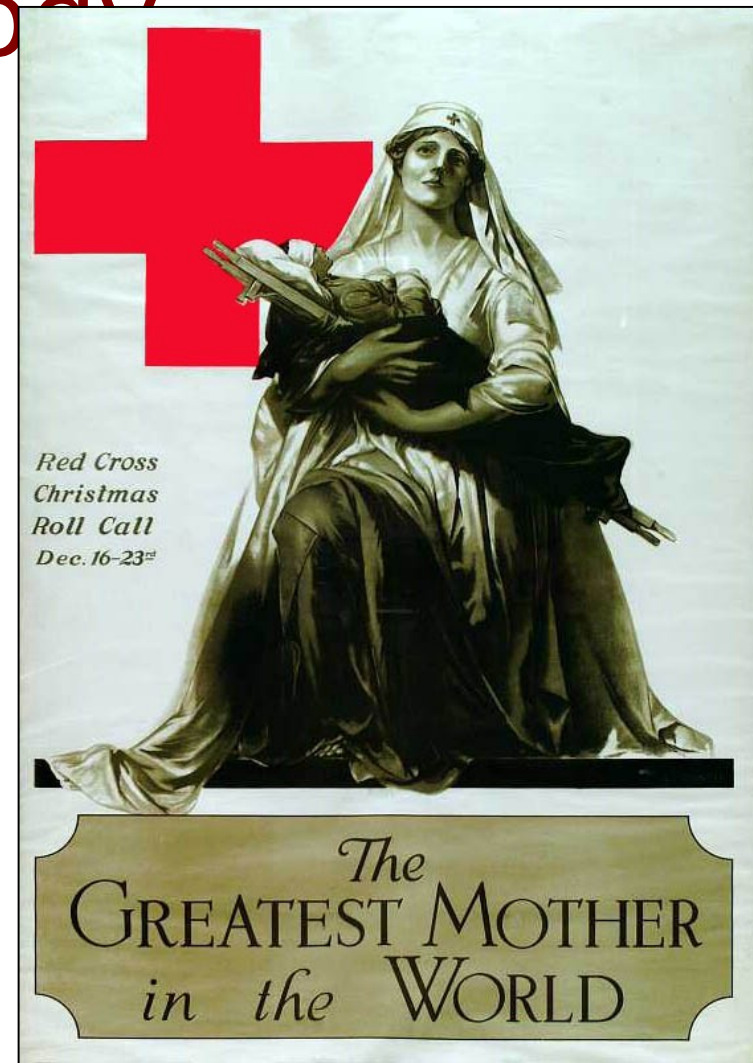
	Head & Neck	Thorax	Abdomen	Limbs
WWI	17	4	2	70
WWII	4	8	4	75
Korean War	17	7	7	74
Vietnam War	14	7	5	74
Gulf War	11	8	7	56
Somalia	20	8	5	65



Combat-related Infections

Bacteriology

- Time of injury
- Echelon III
- Echelon V



Wound Bacteriology

Iraq

- Cultures of US and coalition soldiers
- 31st CSH Baghdad, Iraq
- 49 casualties- 61 wounds

Wound Bacteriology

Pathogens

Gram-positive		Gram-negative	
CNS	32	<i>Pseudomonas stutzeri</i>	1
<i>S. aureus</i>	4 *	<i>C. meningosepticum</i>	1
<i>Micrococcus sp.</i>	1	<i>E. coli</i>	1

Combat-related Infections

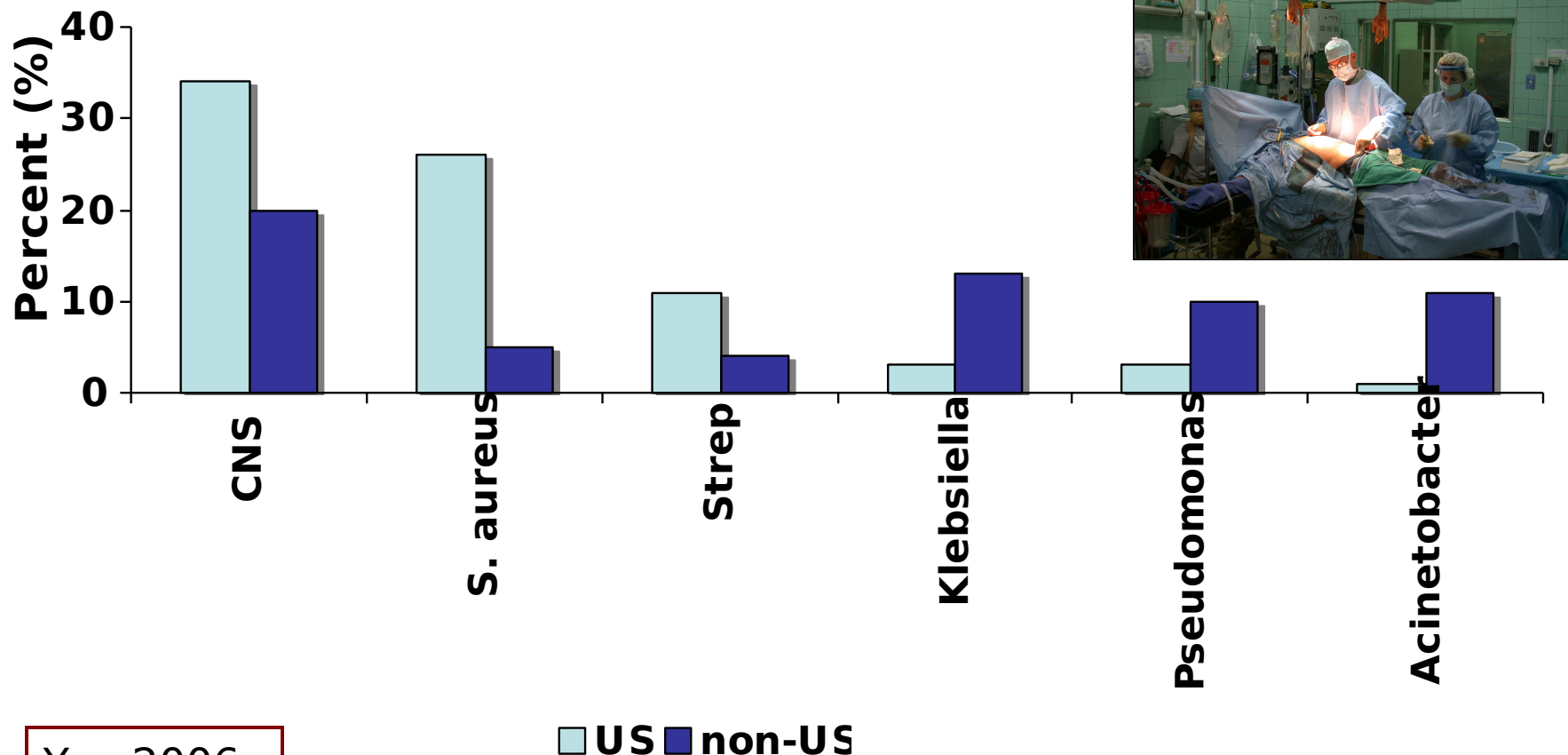
Bacteriology

- Time of injury
- Echelon III
- Echelon V



What Do We Know- In Theater

CSH- Infections 03 04



Yun 2006

What Do We Know- In Theater

Bacteria (#)	1st	2nd	3rd
<i>Acinetobacter</i> (159)	Meropenem (46%)	Amp/sulbac (45%)	Amikacin (41%)
<i>E. coli</i> (93)	Meropenem (100%)	Amikacin (90%)	Pip/tazo (73%)
<i>Klebsiella</i> (66)	Meropenem (95%)	Amikacin (89%)	Gatifloxacin (80%)
<i>Pseudomonas</i> (5)	Amikacin (95%)	*US and non-US personnel (77%)	Pip/tazo (70%)

Provided by Mike Landrum

Wound Bacteriology

USNS Comfort

- March-May 2003
- 300 admissions- 211 trauma patients- 56 infected
 - 85% Iraqi
 - Mean time from injury to admission was 4.2 days

Petersen
2007



Wound Bacteriology

USNS Comfort

- 47 of 56 had wound infections
 - 47% were polymicrobial
- 21 of 56 had blood infection
 - 34% were polymicrobial
- Pathogens
 - *Acinetobacter*- 33%
 - *E. coli*- 14%
 - *Pseudomonas*- 14%

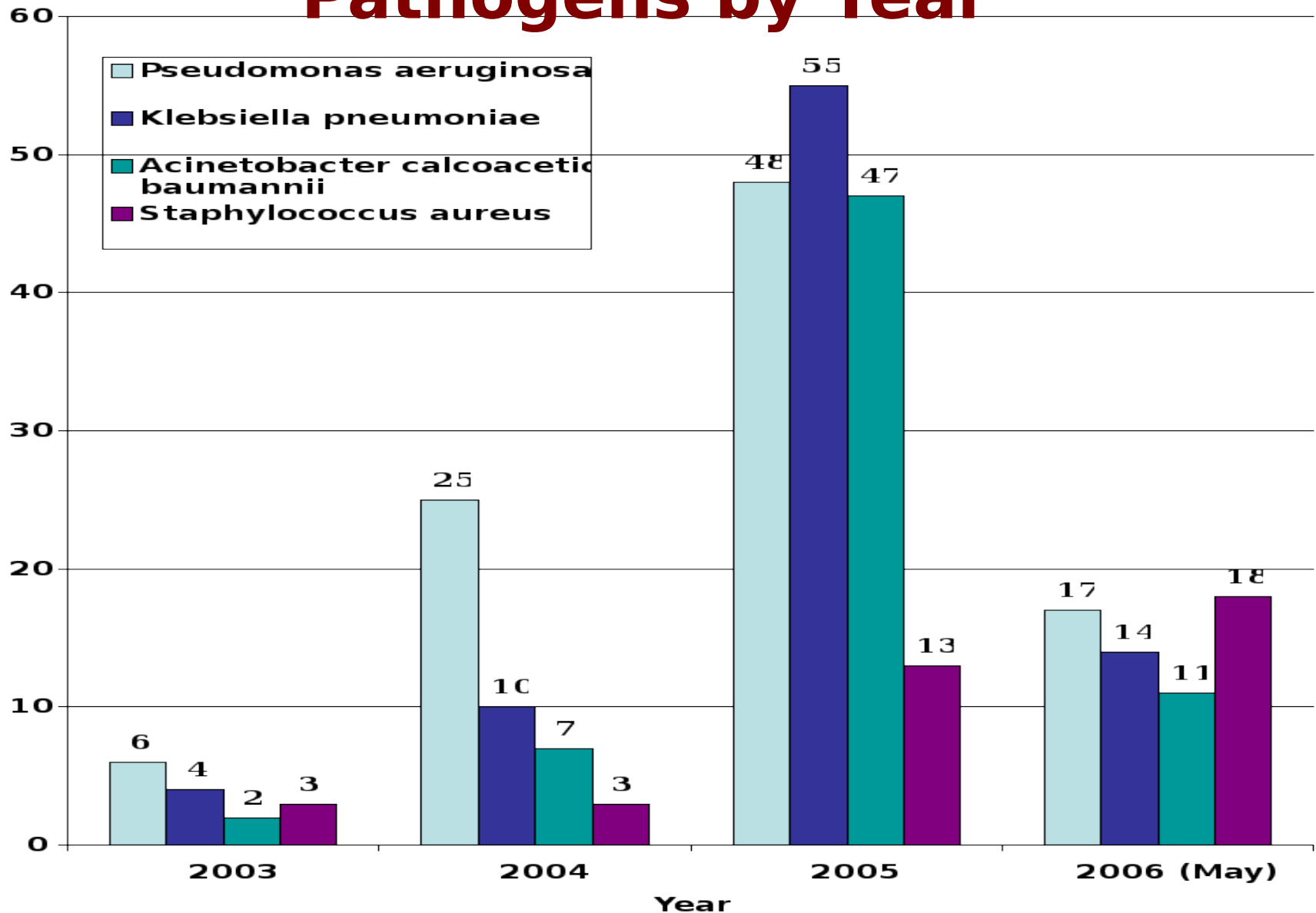
Combat-related Infections

Bacteriology

- Time of injury
- Echelon III
- Echelon V



BAMC Burn ICU- Most Common Pathogens by Year



Combat-Related Infections

Orthopedic Patients

- March 2003-July 2006-
osteomyelitis

- 110 patients with
 - 99 lower extremity
 - 48 upper extremity
 - 2 axial infections



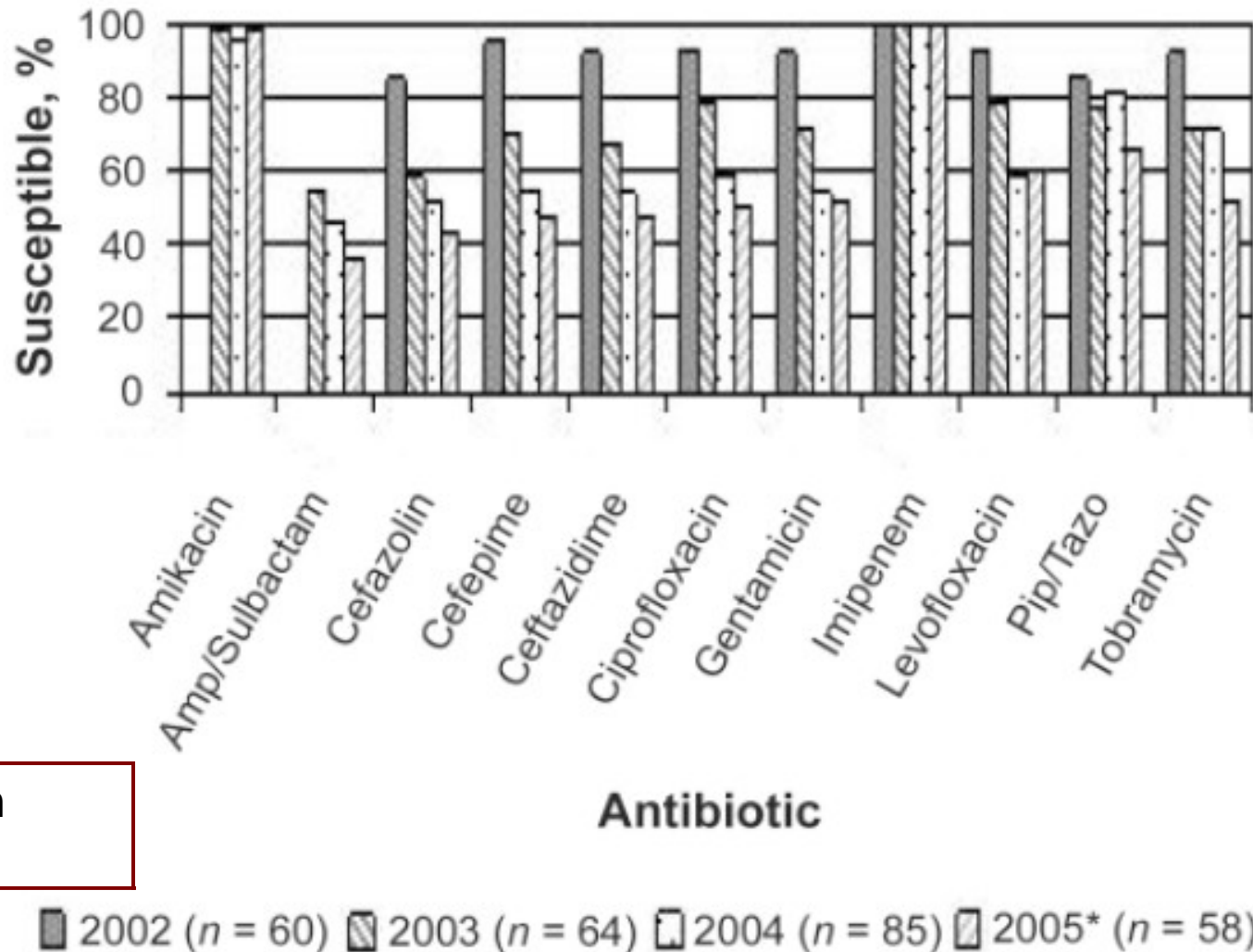
Combat-Related Infections

Orthopedic Patients

- Original versus recurrent/relapse
 - *Acinetobacter* spp.- 71% vs 5%
 - *K. pneumoniae*- 24% vs 5%
 - *P. aeruginosa*- 26% vs 5%
 - *S. aureus*- 15% vs 50%
 - MSSA- 6% vs 22%
 - MRSA- 10% vs 28%

Resistant Bacteria

Klebsiella pneumoniae



Aronson
2006

Combat-related Infections Transmission

- Colonized
- Inoculated at time of injury
- Nosocomial



Resistant Bacteria

Etiology

- Acinetobacter and other MDR pathogens
 - Hospital-acquired infections in Turkey
 - Ventilator-associated pneumonia in Lebanon
 - ICUs in Kuwait
 - Bacteremia in Israel



Resistant Bacteria

Etiology

- Germany
 - 100 a/e patients from Iraq without prior hospital exposure had axilla and groin swabs
 - 0% *Acinetobacter* detected
- Iraq
 - 101 healthy soldiers in Iraq had hands, feet, and head swabs
 - 0% *Acinetobacter* detected



Resistant Bacteria

Etiology

- Inoculated from the environment-
no casualties had gram negative
MDR bacteria



Resistant Bacteria

Etiology

- Nosocomial transmission

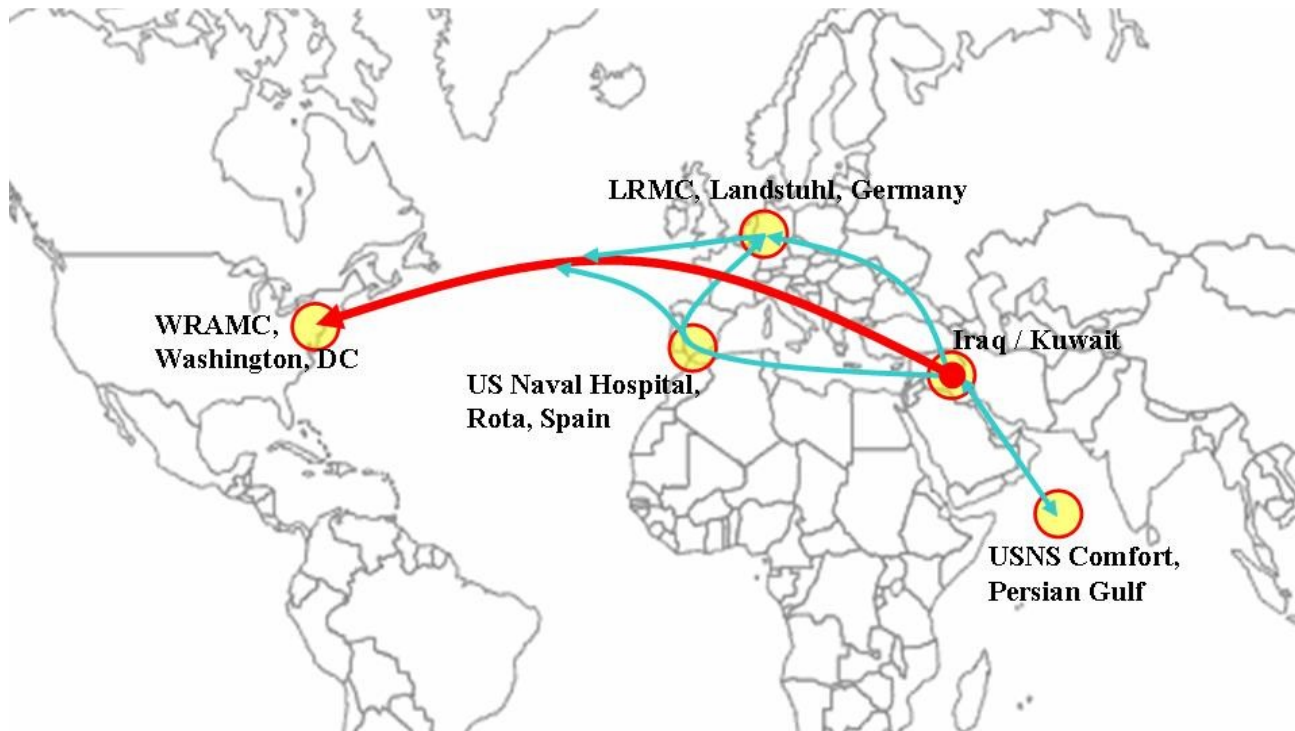


Pictures provided by Stuart Roop

Resistant Bacteria

Etiology

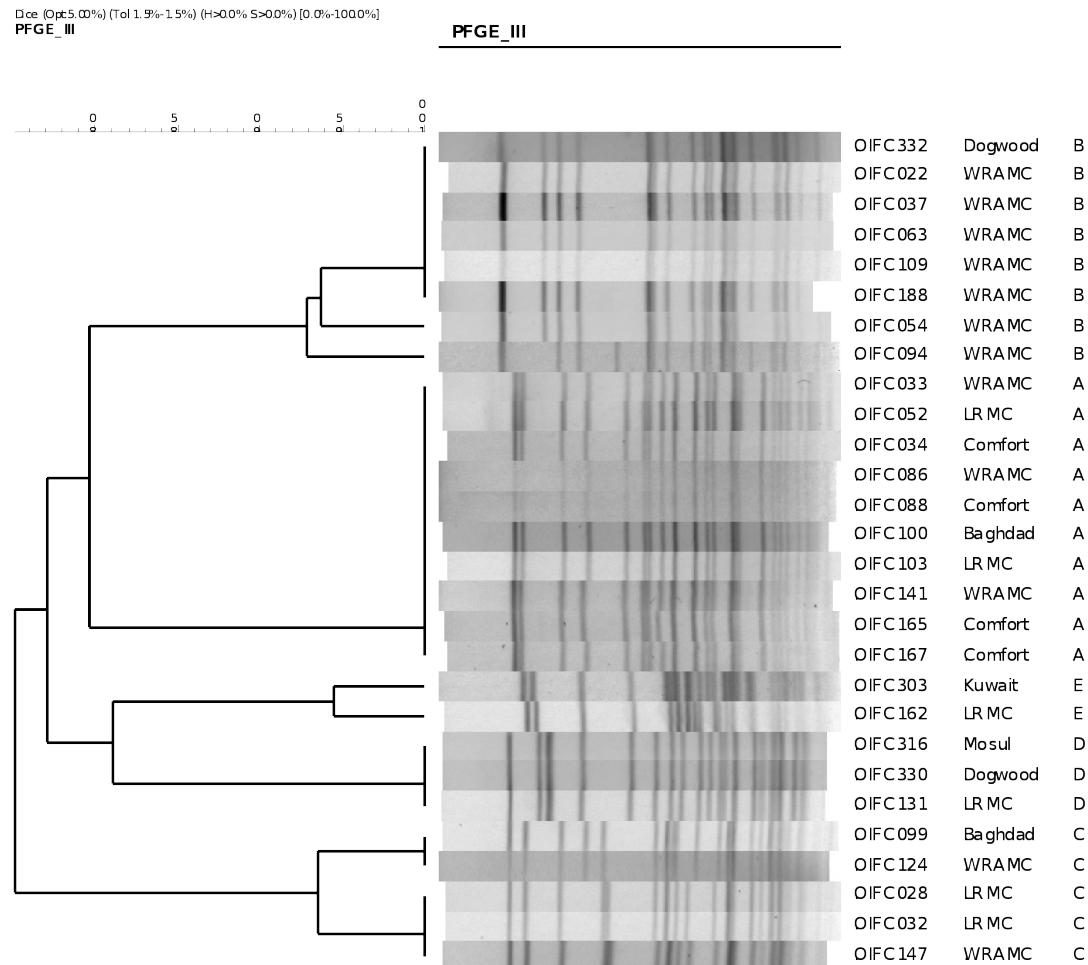
Aeromedical Evacuation Route for Operation Iraqi Freedom (OIF) Casualties, Iraq to WRAMC



Resistant Bacteria

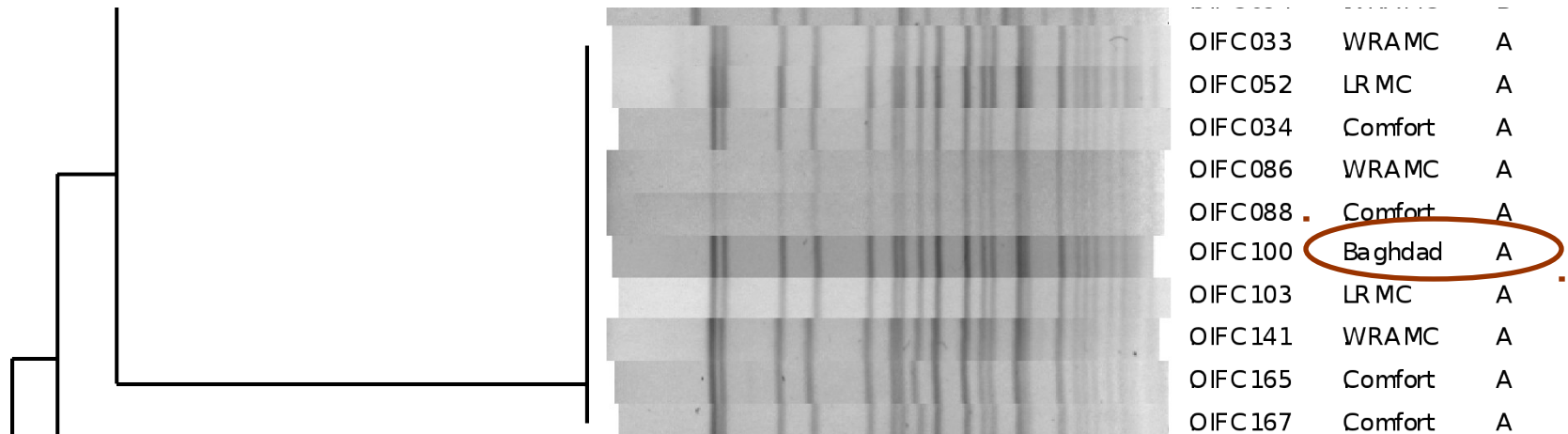
Etiology

- PFGE strains
 - 66 different strains among 170 clinical isolates
 - 25 different strains among 34 environmental isolates



Resistant Bacteria

Etiology



- 43 patients- 2 Baghdad, 18 Comfort, 6 LRMC, 19 WRAMC

Resistant Bacteria

Etiology

- DNA profiles from UK and US isolates identical

United Kingdom and U.S. isolates^d

UK 1	T strain
UK 2	T strain
UK 3	T strain
UK 4	T strain
UK 5	T strain
US 7	T strain
US 9	T strain
US 13	T strain
US 1	T strain
UK 14	OXA-23 clone 2
US 12	OXA-23 clone 2
UK 23	H1AC-2 ^b
UK 10	H1AC-2 ^b
UK 11	H3AC-1 ^b
US 8	USAC-3 ^b

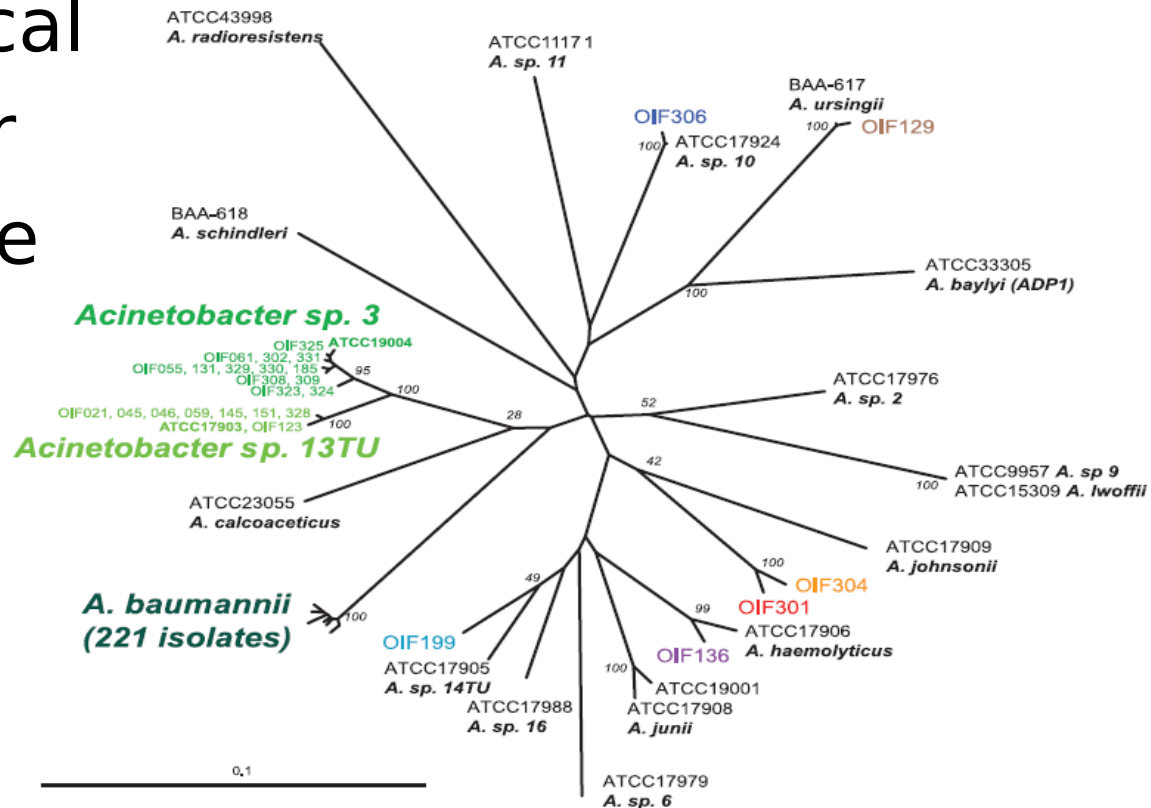
Other United Kingdom isolates

UK 6	W strain
UK 8	W strain
UK 22	V strain
UK 19	SE clone
UK 20	OXA-23 clone 1

Resistant Bacteria

Etiology

- Relatedness to European isolates
 - 14% identical
 - 37% similar
 - 49% diverse



Combat-related Infections

Overview

- Historical review
- Current OIF/OEF epidemiology
- Guideline development
- Guidelines



Combat-related Infections

Working Group

- 11-12 June 2007
 - Triservice with civilians
 - Substantial deployment experience

Combat-related Infections

Assumptions

- Rapid evacuation- injury to US (~7 days)
- Medical facilities- injury to US (~4 sites)
- Varying training and experience



Combat-related Infections

Assumptions

- Some personnel not evacuated



Combat-related Infections

Assumptions

- Multidrug resistant pathogens are infecting war wounded
- No standard prevention guidelines currently exist

Combat-related Infections

Scope- Not Addressed

- Blood transfusion
- Hyperglycemia
- Hypothermia
- Oxygenation



Combat-related Infections

Scope- Not Addressed

- Treatment of nosocomial infections
 - Requires in theater microbiology
 - Requires continually updated antibiogram
 - Rapid de-escalation of antibiotics to monotherapy
 - Minimize peri-operation antibiotics



Combat-related Infections

Target Patient Population

- US, coalition forces- primarily young healthy men without co-morbidities
- Civilian personnel in theater- older with co-morbidities

Combat-related Infections

Target Audience

- Health care providers rendering care to combat-related injuries
- Focused on echelon I-III with echelon IV/V recommendations in supporting manuscripts

Combat-related Infections

Scientific Review

- Experts reviewed literature prior to arrival with emphasis on military studies
- Group discussed all findings/recommendations as a group and then by disseminated of findings/recommendations

Combat-Related Infections

Evidence Based Recommendations

- Strength of Recommendations
 - A- Good evidence to support a recommendation for use
 - B- Moderate evidence to support a recommendation for use
 - C- Poor evidence to support a recommendation for or against use
 - D- Moderate evidence to support recommendation against use
 - E- Good evidence to support a recommendation against use

Combat-Related Infections

Evidence Based Recommendations

- Quality of Evidence
 - I. Evidence from at least one properly randomized controlled trial
 - II. Evidence from at least one well-designed clinical trial without randomization or from cohort or case-controlled studies
 - III. Expert opinion

Combat-related Infections

Overview

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Combat-related Infections

Care at Point of Injury

- Evacuation with surgical evaluation within 6 hours



Combat-related Infections

Care at Point of Injury

- Wounds covered with sterile bandage
- Underlying bony structures stabilized
- If evacuation longer than 3 hours use antibiotics recommended by TCCC
 - Moxifloxacin 400 mg po X 1 OR
 - Ertapenem 1 gm IV/IM X 1

Combat-related Infections

Care without Surgeon- BAS

- Stabilization and evacuation within 6 hours of injury
- Wound irrigation with removal of gross contamination
 - 1-3 L potable water without additives under low pressure



Combat-related Infections

Care without Surgeon- BAS

- Bandage wounds
- Stabilize underlying bony struc



Combat-related Infections

Care without Surgeon- BAS

- Antibiotics
 - Skin, soft tissue, open fractures, exposed bone or open joints
 - Cefazolin 1 gm IV
 - Clindamycin 900 mg IV
 - No enhanced gram negative coverage

Combat-related Infections

Care without Surgeon- BAS

- Antibiotics
 - Abdomen
 - Cefoxitin 1 gm IV
 - Piperacillin-tazobactam- 4.5 gm IV



Combat-related Infections

Care without Surgeon- BAS

- Tetanus toxoid
- Tetanus immunoglobulin
 - No prior immunization and presentation greater than 24 hours
 - Finish tetanus toxoid series



Combat-related Infections

Care without Surgeon- IIa

- Same therapy as BAS
- Retained metal fragment- 1 dose cefazolin
 - Soft tissue injuries only (no fractures, no major vascular involvement and no break of pleura or peritoneum)
 - Wound entry/exit less than 2 cm in maximum dimension
 - Wound not frankly infected
 - Exclusion of mine wounds

Combat-related Infections

Care with Surgeon- IIb/III

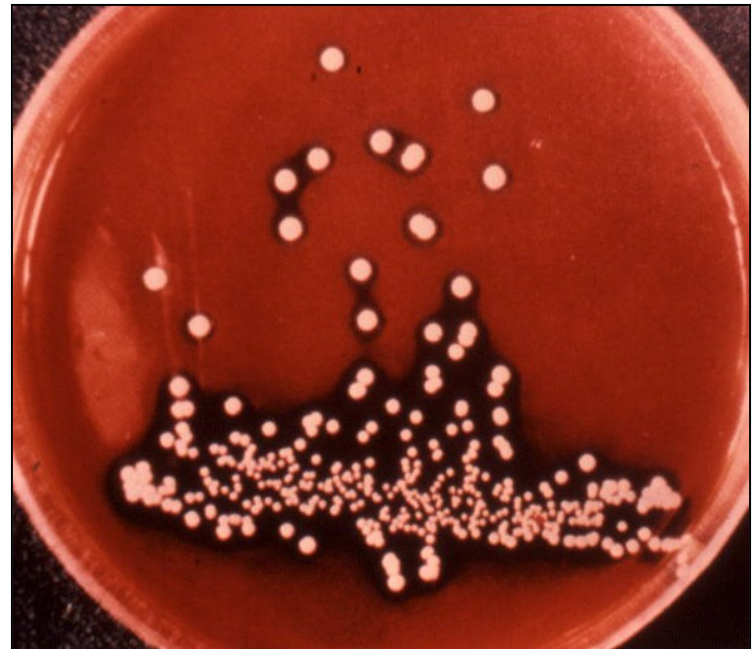
- Surgical evaluation within 6 hours
- Not absolutely necessary for surgical procedure within 6 hours



Combat-related Infections

Care with Surgeon- IIb/III

- No pre/post procedure culture
- Only culture if suspicion of infection

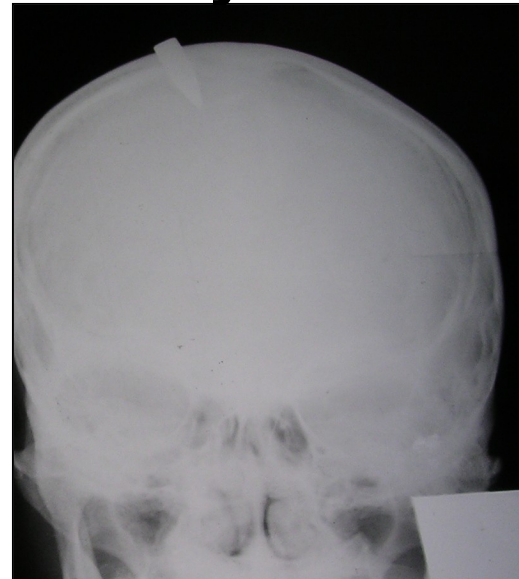


Combat-related Infections

Care with Surgeon- IIb/III

- Aggressive debridement of foreign bodies and necrotic tissue
- Burns debrided within 24 hours
- Delayed removal of foreign body
 - Eye
 - Spine
 - Brain*

* Foreign body may be retained



Combat-related Infections

Care with Surgeon- IIb/III

- Irrigation
 - Bone- Type I fracture- 3L, Type II- 6 L, Type III- 9L
 - Other sites- until contamination removed
 - Fluid- NS or sterile water (tap water ok)
 - No additives
 - Low pressure



Combat-related Infections

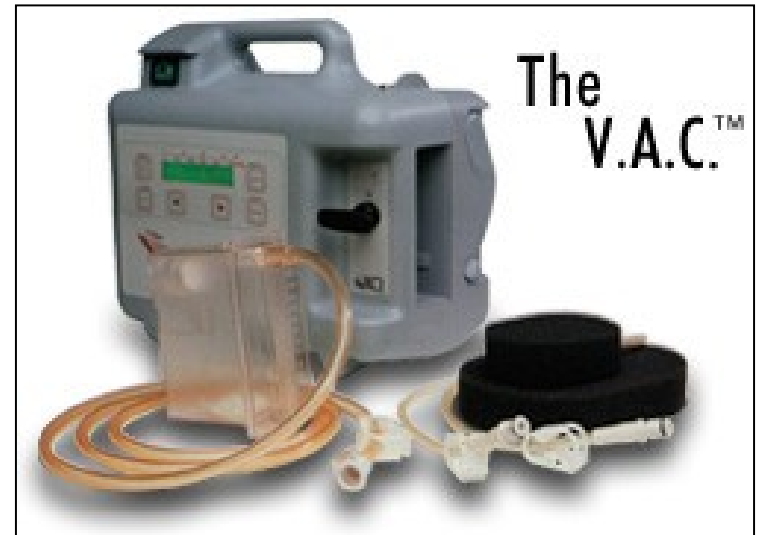
Care with Surgeon- IIb/III

- Antibiotics
 - Avoid broad spectrum agents
 - Short duration of therapy
 - Same as Echelon I/IIa
 - Topical therapy for burn injuries

Combat-related Infections

Care with Surgeon- IIb/III

- Wound closure
 - Delayed primary closure
 - Early closure of face and dura
 - VAC appears effective but concern about air evacuation



Combat-related Infections

Care with Surgeon- IIb/III

- Stabilization of bony structure
 - External fixation appears effective but some concern about infections



Combat-related Infections

Overall

Findings/Recommendation

- Resistant bacteria are complicating our war wounded
- Areas of emphasis
 - Decrease use of broad spectrum antibiotics and prolonged courses
 - Standardized treatment protocols
 - Increase emphasis on basic infection control

Combat-related Infections

Overall

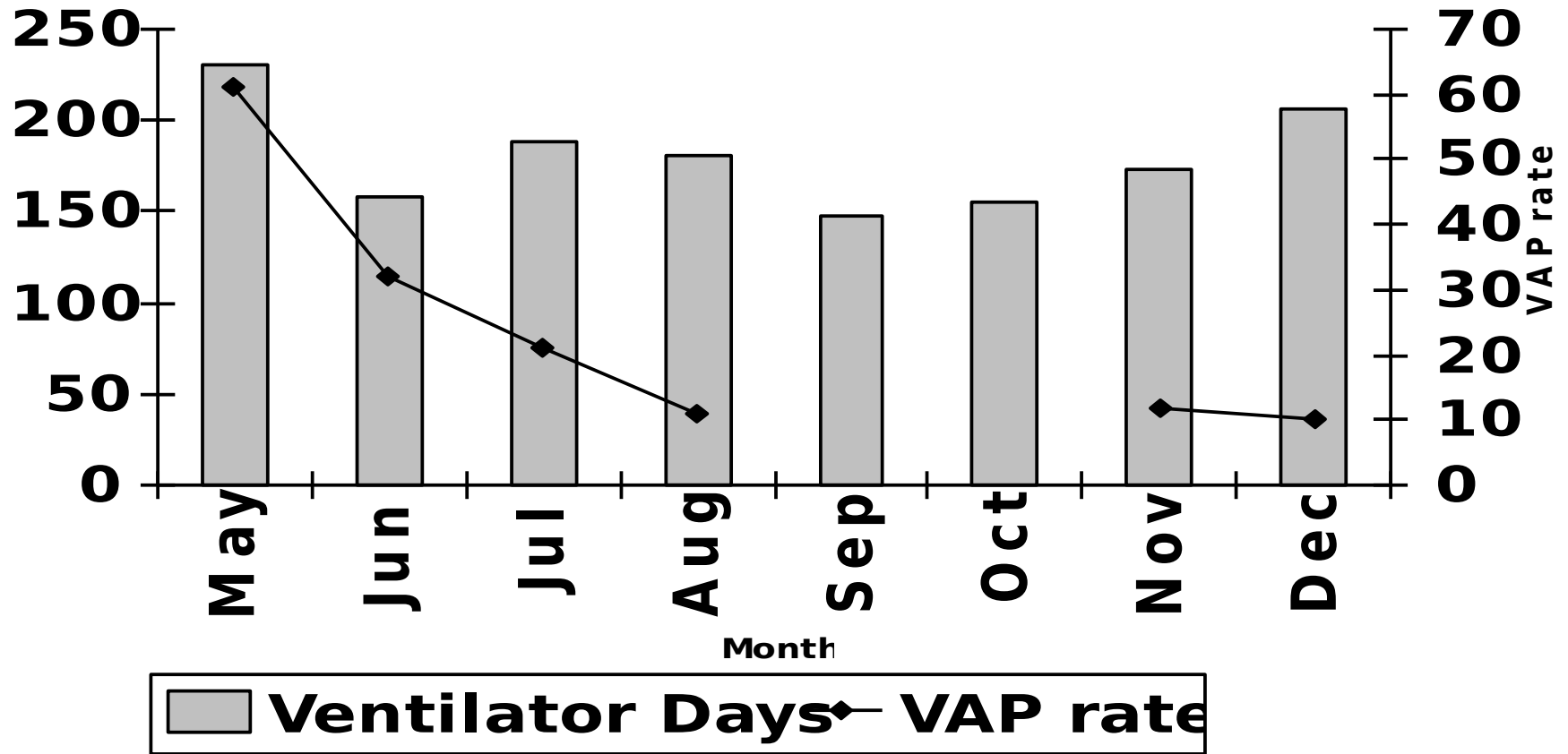
Findings/Recommendation

- Infectious Disease/Infection Control team in theater
 - Antibiotic control programs
 - Hand hygiene
 - Cohorting



Infection Control

VAP Rates

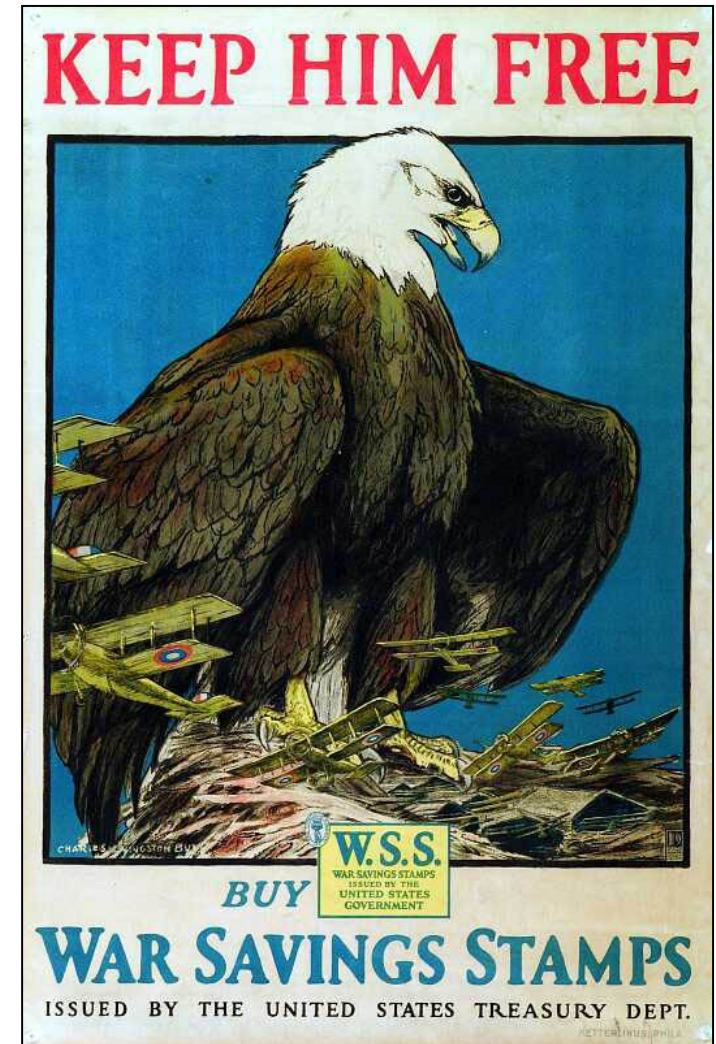


Chi-square for trend,
 $p=0.029$

Combat-related Infections

Overview

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Questions?

